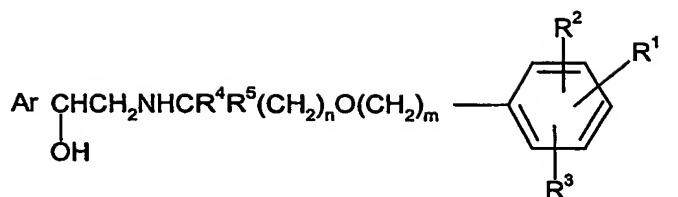


CLAIMS

1. A compound of formula (I)

5



(I)

or a salt, solvate, or physiologically functional derivative thereof, wherein:

10 n is an integer of from 2 to 8;

m is an integer of from 3 to 11, with the proviso that the sum of n + m is from 5 to 19;

R¹ is hydrogen or -XSO₂NR⁶R⁷;

15 wherein X is -(CH₂)_p- or C₂₋₈ alkenylene;

p is an integer from 0 to 6;

R⁶ and R⁷ are independently selected from hydrogen, C₁₋₆alkyl, C₃₋₇cycloalkyl,

20 CONR⁸R⁹, phenyl and phenyl(C₁₋₄alkyl)-,

or R⁶ and R⁷, together with the nitrogen atom to which they are bonded, form a 5-, 6- or 7-membered nitrogen – containing ring;

25 and R⁶ and R⁷ are each independently optionally substituted by 1 or 2 groups independently selected from halo, C₁₋₆alkyl, C₁₋₆alkoxy, hydroxy-substituted C₁₋₆alkoxy, C₁₋₆haloalkyl, CO₂R⁸, SO₂R⁸R⁹, -CONR⁸R⁹, -NR⁸C(O)R⁹ or a 5-, 6- or 7- membered heterocyclic ring;

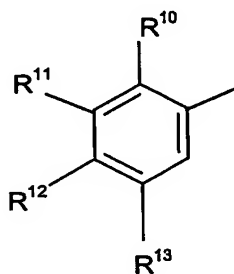
R^8 and R^9 are independently selected from hydrogen, C_{1-6} alkyl, C_{3-7} cycloalkyl, phenyl and phenyl(C_{1-6} alkyl)-;

5 R^2 and R^3 are independently selected from hydrogen, C_{1-6} alkyl, C_{1-6} alkoxy, halo, phenyl and C_{1-6} haloalkyl;

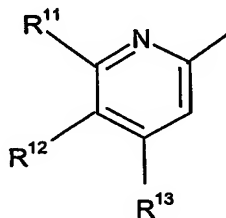
R^4 and R^5 are independently selected from hydrogen and C_{1-4} alkyl with the proviso that the total number of carbon atoms in R^4 and R^5 is not more than 4,

10 and

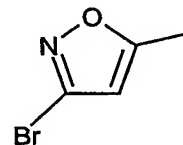
Ar is a group selected from



(a)

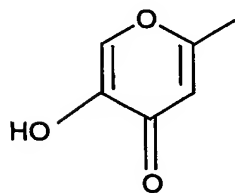


(b)



(c)

and



(d)

15

wherein R^{11} represents hydrogen, halogen, $-(CH_2)_4OR^{14}$, $-NR^{14}C(O)R^{15}$, $-NR^{14}SO_2R^{15}$, $-SO_2NR^{14}R^{15}$, $-NR^{14}R^{15}$, $-OC(O)R^{16}$ or $OC(O)NR^{14}R^{15}$,

and R^{10} represents hydrogen, halogen or C_{1-4} alkyl;

or R^{11} represents $-NHR^{17}$ and R^{10} and $-NHR^{17}$ together form a 5- or 6- membered heterocyclic ring;

5

R^{12} represents hydrogen, halogen, $-OR^{14}$ or $-NR^{14}R^{15}$; $-OC(O)R^{16}$ or $-OC(O)NR^{14}R^{15}$;

R^{13} represents hydrogen, halogen, halo C_{1-4} alkyl, $-OR^{14}$ or $-NR^{14}R^{15}$;

10 R^{14} and R^{15} each independently represents hydrogen or C_{1-4} alkyl, or in the groups $-NR^{14}R^{15}$, $-SO_2NR^{14}R^{15}$ and $-OC(O)NR^{14}R^{15}$, R^{14} and R^{15} independently represent hydrogen or C_{1-4} alkyl or together with the nitrogen atom to which they are attached form a 5-, 6- or 7- membered nitrogen-containing ring,

15 R^{16} represents an aryl (eg phenyl or naphthyl) group which may be unsubstituted or substituted by one or more substituents selected from halogen, C_{1-4} alkyl, hydroxy, C_{1-4} alkoxy or halo C_{1-4} alkyl; and

q is zero or an integer from 1 to 4;

20

provided that when R^1 is hydrogen

Ar is not a group (a) wherein;

R^{11} is $-(CH_2)_qOR^{14}$, q is zero or 1 and R^{12} is OR^{14} ,

or R^{11} is $-(CH_2)_qOR^{14}$, q is zero and R^{13} is OR^{14} ,

25 or R^{11} is $-NR^{14}SO_2R^{15}$ or $NR^{14}COR^{15}$ and R^{12} is OR^{14} ,

or R^{11} and R^{13} both represent halogen and R^{12} is $NR^{14}R^{15}$;

Ar is not a group (b) wherein R^{11} is $-(CH_2)_qOR^{14}$ and R^{12} is OR^{14} ;

Ar is not a group (c),

and when R^1 is $XSO_2NR^6R^7$, Ar is not a group (a) wherein

30 R^{11} is $(CH_2)_qOR^{14}$ or $NR^{14}COR^{15}$, and R^{12} is OR^{14} .

2. A compound of formula (I) according to claim 1 wherein, in the group Ar, R^{11} represents halogen, $-(CH_2)_qOR^{14}$, $-NR^{14}C(O)R^{15}$, $-NR^{14}SO_2R^{15}$, $-SO_2NR^{14}R^{15}$, $-NR^{14}R^{15}$, $-OC(O)R^{16}$ or $OC(O)NR^{14}R^{15}$,

35

and R¹⁰ represents hydrogen,

or R¹¹ represents -NHR¹⁷ and R¹⁰ and -NHR¹⁷ together form a 5- or 6- membered heterocyclic ring;

5

and

R¹³ represents hydrogen, halogen, haloC₁₋₄ alkyl, -OR¹⁴, or -NR¹⁴R¹⁵;

10 and all other substituents are as defined in claim 1.

3. A compound of formula (I) according to claim 1 or claim 2 wherein the group R¹ is attached to the meta-position relative to the -O-(CH₂)_m link.

15 4. A compound of formula (I) according to any of claims 1 to 3 wherein R¹ represents SO₂NR⁶R⁷ wherein R⁶ and R⁷ are independently selected from hydrogen and C₁₋₆alkyl.

5. A compound of formula (I) according to any of claims 1 to 4 wherein R⁴ and R⁵ are independently selected from hydrogen and methyl.

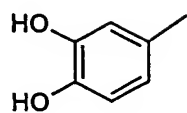
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6. A compound of formula (I) according to any of claims 1 to 5 wherein R² and R³ each represent hydrogen.

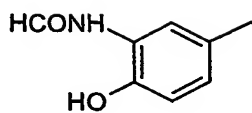
7. A compound of formula (I) according to any of claims 1 to 6 wherein n is 5 or 6 and
25 m is 3 or 4 such that m + n is 8, 9 or 10.

8. A compound of formula (I) according to any of claims 1 to 7 wherein Ar represents a group selected from:

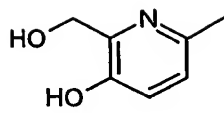
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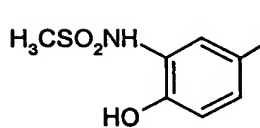
(i)



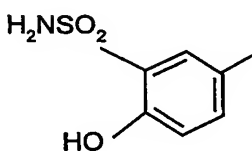
(ii)



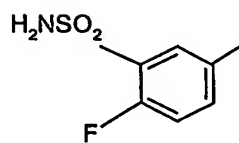
(iii)



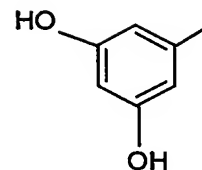
(iv)



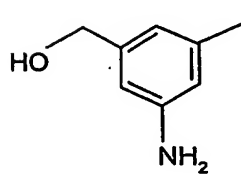
(v)



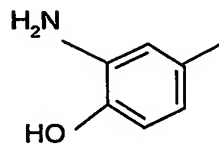
(vi)



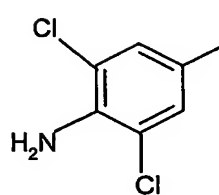
(vii)



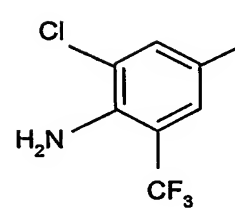
(viii)



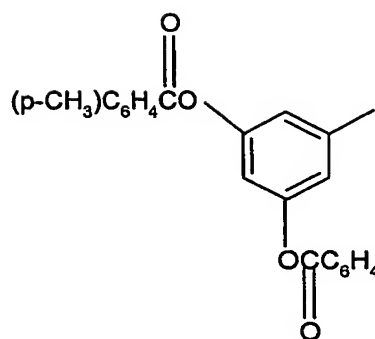
(ix)



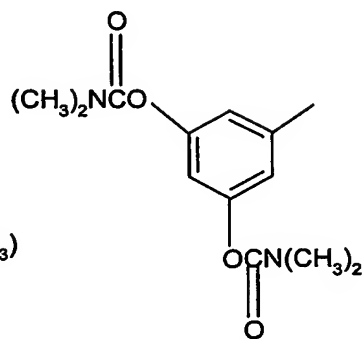
(x)



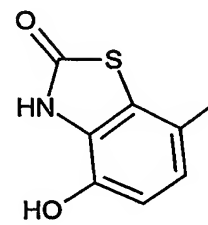
(xi)



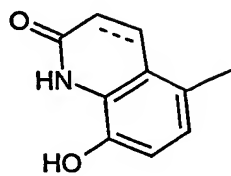
(xii)



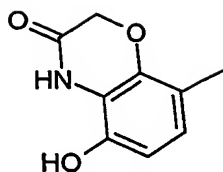
(xiii)



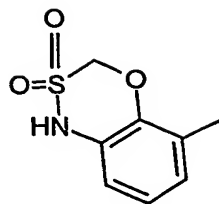
(xiv)



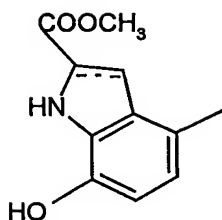
(xv)



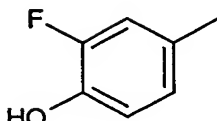
(xvi)



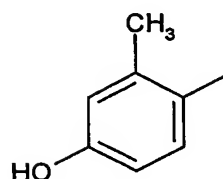
(xvii)



(xviii)



(xix)



(xx)

5 9. A compound of formula (I) according to any of claims 1 to 8 wherein R^1 is hydrogen and Ar is selected from a group of structure (ii), (v), (vi), (viii), (ix), (xi), (xii), (xiii), (xiv), (xv), (xvi), (xvii) and (xviii).

10 10. A compound of formula (I) according to any of claims 1 to 8 wherein R^1 is $XSO_2NR^6R^7$ and Ar is selected from a group of structure (iii), (iv), (xiv), (xv), (xvi) and (xix)

11. A compound of formula (I) selected from:
 8-Hydroxy-5-((1*R*)-1-hydroxy-2-([6-(4-phenylbutoxy)hexyl]amino)ethyl)quinolin-2(1*H*)-one;
 3-{4-[(6-[(2*R*)-2-Hydroxy-2-(8-hydroxy-2-oxo-1,2-dihydroquinolin-5-yl)ethyl]amino)hexyl]oxy]butyl}benzenesulfonamide;
 5-Hydroxy-8-(1-hydroxy-2-([6-(4-phenylbutoxy)hexyl]amino)ethyl)-2*H*-1,4-benzoxazin-3(4*H*)-one;
 3-{4-[(6-[[2-hydroxy-2-(5-hydroxy-3-oxo-3,4-dihydro-2*H*-1,4-benzoxazin-8-yl)ethyl]amino]hexyl]oxy]butyl}benzenesulfonamide;

4-Hydroxy-7-((1*R*)-1-hydroxy-2-[[6-(4-phenylbutoxy)hexyl]amino]ethyl)-1,3-benzothiazol-2(3*H*)-one;

4-Hydroxy-7-(1-hydroxy-2-[[6-(4-phenylbutoxy)hexyl]amino]ethyl)-1,3-benzothiazol-2(3*H*)-one;

- 5 3-{4-[[6-[(2*R*)-2-(3-Fluoro-4-hydroxyphenyl)-2-hydroxyethyl]amino]hexyl]oxy}butyl]benzenesulfonamide;
- 3-(4-[[6-[(2-Hydroxy-2-[5-hydroxy-6-(hydroxymethyl)pyridin-2-yl]ethyl]amino)hexyl]oxy}butyl]benzenesulfonamide;
- 3-[4-[(6-[(2*R*)-2-Hydroxy-2-[4-hydroxy-3-
10 [(methylsulfonyl)amino]phenyl)ethyl]amino]hexyl]oxy)butyl]benzenesulfonamide;
- 3-{3-[(7-[(2*R*)-2-(3-Fluoro-4-hydroxyphenyl)-2-hydroxyethyl]amino)heptyl]oxy}propyl]benzenesulfonamide;
- 3-(3-[(7-[(2-Hydroxy-2-[5-hydroxy-6-(hydroxymethyl)pyridin-2-yl]ethyl]amino)heptyl]oxy}propyl]benzenesulfonamide;
- 15 3-[3-[(7-[(2*R*)-2-Hydroxy-2-[4-hydroxy-3-[(methylsulfonyl)amino]phenyl)ethyl]amino]heptyl]oxy}propyl]benzenesulfonamide;
- 3-{3-[(7-[(2*R*)-2-Hydroxy-2-(8-hydroxy-2-oxo-1,2-dihydroquinolin-5-yl)ethyl]amino)heptyl]oxy}propyl]benzenesulfonamide;
- 3-(3-[(7-[(2*R*)-2-[3-(Formylamino)-4-hydroxyphenyl]-2-
20 hydroxyethyl]amino)heptyl]oxy}propyl]benzenesulfonamide;

or a salt, solvate or physiologically functional derivative thereof.

12. A method for the prophylaxis or treatment of a clinical condition in a mammal, such
25 as a human, for which a selective β_2 -adrenoreceptor agonist is indicated, which comprises administration of a therapeutically effective amount of a compound of formula (I) according to any of claims 1 to 11, or a pharmaceutically acceptable salt, solvate, or physiologically functional derivative thereof.

- 30 13. A compound of formula (I), according to any of claims 1 to 11, or a pharmaceutically acceptable salt, solvate, or physiologically functional derivative thereof for use in medical therapy.

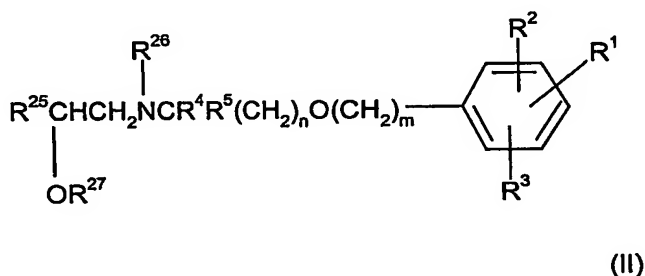
14. A pharmaceutical formulation comprising a compound of formula (I), according to
35 any of claims 1 to 11, or a pharmaceutically acceptable salt, solvate, or physiologically

functional derivative thereof, and a pharmaceutically acceptable carrier or excipient, and optionally one or more other therapeutic ingredients.

15. The use of a compound of formula (I), according to any of claims 1 to 11, or a pharmaceutically acceptable salt, solvate, or physiologically functional derivative thereof in the manufacture of a medicament for the prophylaxis or treatment of a clinical condition for which a selective β_2 -adrenoreceptor agonist is indicated.

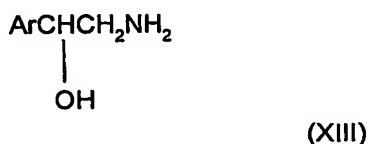
16. A process for the preparation of a compound of formula (I), according to any of claims 1 to 11, or a salt, solvate, or physiologically functional derivative thereof, which comprises:

(a) deprotection of a protected intermediate of formula (II):

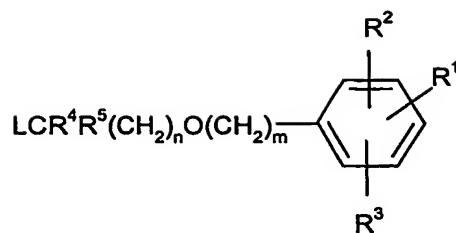


or a salt or solvate thereof, wherein R^1 , R^2 , R^3 , R^4 , R^5 , m and n are as defined for the compounds of formula (I) R^{25} represents an optionally protected form of Ar, and R^{26} and R^{27} each independently represent either hydrogen or a protecting group, provided that the compound of formula (II) contains at least one protecting group;

(b) reacting a compound of formula (XIII):



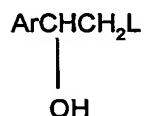
Wherein Ar is as defined above with a compound of formula (VI):



(VI)

Wherein L is a leaving group such as halo (typically chloro, bromo or iodo) or a sulphonate (typically methanesulphonate) and R^1 , R^2 , R^3 , R^4 , R^5 , n and m are as defined
 5 for compounds of formula (I).

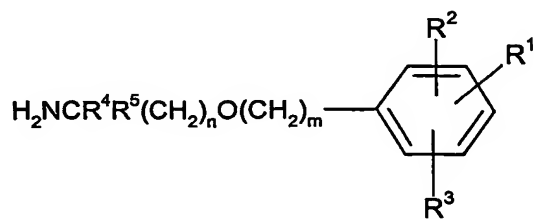
(c) reacting a compound of formula (XV):



(XV)

10

wherein L is a leaving group as hereinbefore defined, with an amine of formula (XVI):



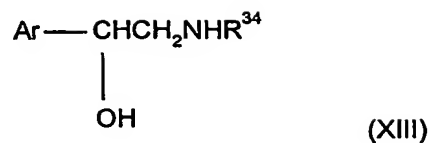
(XVI)

15

wherein R^1 , R^2 , R^3 , R^4 , R^5 , n and m are as defined for formula (I); and

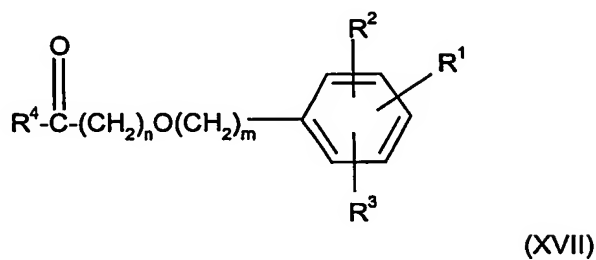
(d) (i) reacting a compound of formula (XIII):

20



Wherein Ar is as hereinbefore defined and R^{34} is a chiral auxiliary group,

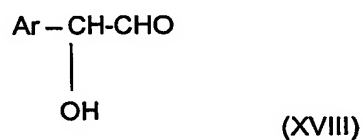
5 with a compound of formula (XVII):



wherein R^1 , R^2 , R^3 , R^4 , n and m are as hereinbefore defined;
followed where necessary by removal of said chiral auxiliary group R^{34} ;

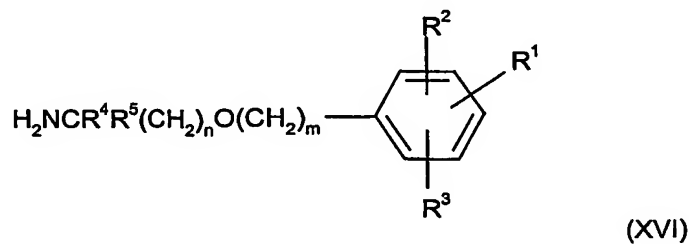
10

or (ii) reacting a compound of formula (XVIII):



wherein Ar is as hereinbefore defined; with an amine of formula (XVI):

15



as hereinbefore defined,

5 under conditions suitable to effect reductive amination,

followed by the following steps in any order:

- 10 (i) optional removal of any protecting groups;
(ii) optional separation of an enantiomer from a mixture of enantiomers;
(iii) optional conversion of the product to a corresponding salt, solvate,
(iv) optional conversion of a group R^1 , R^2 and/or R^3 to another group R^1 , R^2
and/or R^3 ,

or physiologically functional derivative thereof.

15

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